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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
10/082,772	02/25/2002	Peter Droge	DEBE:008US	4391	
7590 02/27/2007 Steven L. Highlander FULBRIGHT & JAWORSKI L.L.P.			EXAM	EXAMINER	
			NGUYEN, QUANG		
Suite 2400 600 Congress Avenue,		ART UNIT	PAPER NUMBER		
Austin, TX 787		•	1633		
			MAIL DATE	DELIVERY MODE	
			02/27/2007	PAPER	

Please find below and/or attached an Office communication concerning this application or proceeding.

	Application No.	Applicant(s)			
Advisory Action	10/082,772	DROGE ET AL.			
Before the Filing of an Appeal Brief	Examiner	Art Unit			
	Quang Nguyen, Ph.D.	1633			
The MAILING DATE of this communication appears on the cover sheet with the correspondence address					
THE REPLY FILED 14 February 2007 FAILS TO PLACE THIS	APPLICATION IN CONDITION FO	R ALLOWANCE.			
 The reply was filed after a final rejection, but prior to or of this application, applicant must timely file one of the folk places the application in condition for allowance; (2) a Na Request for Continued Examination (RCE) in compliant time periods: The period for reply expiresmonths from the mailing the period of t	owing replies: (1) an amendment, affortice of Appeal (with appeal fee) in once with 37 CFR 1.114. The reply ming date of the final rejection.	idavit, or other evider compliance with 37 C ust be filed within one	nce, which FR 41.31; or (3) of the following		
b) The period for reply expires on: (1) the mailing date of this no event, however, will the statutory period for reply expire Examiner Note: If box 1 is checked, check either box (a) of TWO MONTHS OF THE FINAL REJECTION. See MPEP	later than SIX MONTHS from the mailing (b). ONLY CHECK BOX (b) WHEN THE 706.07(f).	g date of the final rejecting E FIRST REPLY WAS F	on. ILED WITHIN		
Extensions of time may be obtained under 37 CFR 1.136(a). The dat have been filed is the date for purposes of determining the period of e under 37 CFR 1.17(a) is calculated from: (1) the expiration date of the set forth in (b) above, if checked. Any reply received by the Office lat may reduce any earned patent term adjustment. See 37 CFR 1.704(INOTICE OF APPEAL	xtension and the corresponding amount shortened statutory period for reply origer than three months after the mailing day).	of the fee. The appropri inally set in the final Offi te of the final rejection, o	iate extension fee ce action; or (2) as even if timely filed,		
2. The Notice of Appeal was filed on A brief in compliance with 37 CFR 41.37 must be filed within two months of the date of filing the Notice of Appeal (37 CFR 41.37(a)), or any extension thereof (37 CFR 41.37(e)), to avoid dismissal of the appeal. Since a Notice of Appeal has been filed, any reply must be filed within the time period set forth in 37 CFR 41.37(a). AMENDMENTS					
3. The proposed amendment(s) filed after a final rejection (a) They raise new issues that would require further c (b) They raise the issue of new matter (see NOTE bel (c) They are not deemed to place the application in be appeal; and/or (d) They present additional claims without canceling a	onsideration and/or search (see NO ow); etter form for appeal by materially re	TE below);	i		
NOTE: See Continuation Sheet. (See 37 CFR 1.116 and 41.33(a)).					
4. The amendments are not in compliance with 37 CFR 1.121. See attached Notice of Non-Compliant Amendment (PTOL-324). 5. Applicant's reply has overcome the following rejection(s):					
 Newly proposed or amended claim(s) would be a non-allowable claim(s). 		timely filed amendme	ent canceling the		
7. For purposes of appeal, the proposed amendment(s): a how the new or amended claims would be rejected is profile that the status of the claim(s) is (or will be) as follows: Claim(s) allowed: Claim(s) objected to: Claim(s) rejected: 29,30,32-39,43-51 and 58. Claim(s) withdrawn from consideration:)⊠ will not be entered, or b) □ wil ovided below or appended.	ll be entered and an e	explanation of		
AFFIDAVIT OR OTHER EVIDENCE					
 The affidavit or other evidence filed after a final action, because applicant failed to provide a showing of good at was not earlier presented. See 37 CFR 1.116(e). 	nd sufficient reasons why the affidav	it or other evidence is	necessary and		
 The affidavit or other evidence filed after the date of filing entered because the affidavit or other evidence failed to showing a good and sufficient reasons why it is necessangled. The affidavit or other evidence is entered. An explanation of the evidence is entered. 	overcome <u>all</u> rejections under appeary and was not earlier presented. S	al and/or appellant fai ee 37 CFR 41.33(d)(1	ls to provide a		
REQUEST FOR RECONSIDERATION/OTHER		•			
11. The request for reconsideration has been considered b See Continuation Sheet.		o condition for allowar	nce Decause:		
12. ☐ Note the attached Information Disclosure Statement(s).13. ☐ Other:	(P10/SB/08) Paper No(s)				
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Continuation of 3. NOTE: The scope of the proposed claims is not the same as that of finally rejected claims, which would require further consideration and search given the newly recited limitation "a first DNA segment stably intrgrated into the genome of said cell".

Continuation of 11. does NOT place the application in condition for allowance because: Applicant's arguments are respectfully not found persuasive for overcoming the prior art of record. It is noted that Applicants presented the same arguments as those in the amendment filed on 10/4/06, and these arguments have been addressed in details in the Final office action mailed on 12/14/06. With respect to any argument related to the proposed claims, it is moot because the proposed claims were not entered. With respect to the data of Lange-Gustafson et al., Applicants argue that the studies were performed in vitro and reflect conditions (25 C and KCI at 25 mM) that have nothing to do with the environment inside a living eukarytotic cell.. Lange-Gustafson taught clearly that Int-h uses supercoiled DNA more effectively than non-supercoiled DNA as a substrate for recombination, and that in the absence of IHF, Int-h recombines supercoiled and nonsupercoiled substrates identically (see abstract) at least in vitro. There is no factual evidence that Int-h would not be able to use supercoiled DNA at all under physiological conditions or inside a eukaryotic cell; and Applicants do not provide any factual evidence to support Applicants' argument. Furthermore, as already noted in the Final office action since wild-type lambda integrases are capable of mediating sequence specific recombination events in both eukaryotic and prokaryotic cells, and it is known that the conditions required by a wild-type lambda integrase to mediate a sequence specific recombination event in prokaryotic cells, under physiological conditions and in vitro conditions are even more stringent than those required by the Int-h (see the teachings of Hartley et al., Christ & Droge and Lange-Gustafson), it is therefore reasonable for an ordinary skilled artisan to expect that at least Int-h is also able to function in eukaryotic cells to mediate sequence specific recombination of both eukaryotic and/or prokaryotic DNA segments.

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